BRIEF REPORT

Cross-Sectional Validation of Diabetes Risk Scores for Predicting Diabetes, Metabolic Syndrome, and Chronic Kidney Disease in Taiwanese

Jou-Wei Lin, md, phd^{1,2} Yi-Cheng Chang, md¹ Hung-Yuan Li, md, ms^{2,3} Yu-Fen Chien, ms¹ Mei-Yu Wu, rn¹ Ru-Yi Tsai, rn¹ Yenh-Chen Hsieh, bs¹ Yu-Jen Chen, bs¹ Juey-Jen Hwang, md, phd^{2,3} Lee-Ming Chuang, md, phd^{2,3}

OBJECTIVE — To validate the performance of current diabetes risk scores (DRSs) based on simple clinical information in detecting type 2 diabetes, metabolic syndrome (MetSyn), and chronic kidney disease (CKD).

RESEARCH DESIGN AND METHODS — The performance of 10 DRSs was evaluated in a cross-sectional population screening of 2,759 Taiwanese subjects.

RESULTS — All DRSs significantly correlated with measures of insulin resistance, estimated glomerular filtration rate, and urine albumin excretion. The prevalence of screening-detected diabetes (SDM), MetSyn, and CKD increased with higher DRSs. For prediction of SDM, the Cambridge DRS by Griffin et al. and the Finnish DRS outperformed other DRSs in terms of discriminative power and model fit. For prediction of MetSyn and CKD, the Atherosclerosis Risk in Community Study score by Schmidt et al. outperformed other DRSs.

CONCLUSIONS — Risk scores based on simple clinical information are useful to identify individuals at high risk for diabetes, MetSyn, and CKD in different ethnic populations.

Diabetes Care 32:2294-2296, 2009

he use of fasting or postchallenge plasma glucose concentrations has been proposed for the early identification of type 2 diabetes. However, both tests are costly and time-consuming. Therefore, a simple diabetes risk score (DRS) that does not require any laboratory test is needed to identify individuals at high risk. This study aimed to systematically evaluate the performance of current DRSs based on simple clinical information in identifying diabetes in a cross-sectional population screening in Taiwanese. The feasibility of these scores in identifying individuals at high risk for

metabolic syndrome (MetSyn) and chronic kidney disease (CKD), two conditions closely related to type 2 diabetes, were also evaluated.

RESEARCH DESIGN AND METHODS

From 8 June 2005 to 22 November 2008, 2,759 participants undergoing community-based screening for type 2 diabetes in the Yun-Lin area in Taiwan were recruited. The exclusion criteria were age <18 years, pregnant women, previously diagnosed diabetes, or previously diagnosed renal disease. Measures of insulin resis-

tance and β-cell function were determined using the homeostasis model assessment (HOMA)-2 with the use of a HOMA calculator (www.dtu.ox.ac.uk). The glomerular filtration rate (GFR) was estimated by the Modification of Diet in Renal Disease (MDRD) Study equation (1). Urine albumin excretion was calculated using a random urine albumin-tocreatinine ratio. Type 2 diabetes was diagnosed by a fasting plasma glucose concentration >126 mg/dl. MetSyn was defined according to the National Cholesterol Education Program Third Adult Treatment Panel guideline with modification for Asian populations (2). CKD was defined as MDRD GFR < 60 ml/min per $1.73 \,\mathrm{m}^2$ (1). The cutoff values for BMI (24) and 27 kg/m²) and waist circumference (90 cm for men; 80 cm for women) were modified according to the definition of obesity for Taiwanese and the modified waist circumference criteria for Asians (2). Ten currently available DRSs, including the Atherosclerosis Risk in Community (ARIC) Study, Asian Indian, Cambridge (U.K.), Danish, DESIR (Data from the Epidemiological Study on Insulin Resistance Syndrome) (French), Dutch, FINDRISC (Finnish diabetes risk score), Oman, QDScore (U.K.), and Thai risk scores, were analyzed (3-12). The institutional review board approved this study, and informed consent was obtained from each participant. The area under the receiver operating characteristic (ROC) curves was used to assess the discriminative power of DRSs. Model fitness was assessed by the Hosmer-Lemeshow test, and the DeLong method was used to compare areas under the ROC curves.

RESULTS

All DRSs correlated significantly with components of MetSyn, HOMA2–insulin resistance, high-sensitive C-reactive protein, and uric acid levels, all of which were markers of insulin resistance (supplementary Table S1, found in an online appendix at http://care.diabetesjournals.org/cgi/content/full/dc09-0694/DC1). However, HOMA2- β , a measure of insulin secretion, did not

From the ¹Metabolic Syndrome and Diabetes Screening Program, National Taiwan University Hospital Yun-Lin Branch, Dou-Liou City, Taiwan; the ²Department of Medicine, College of Medicine, National Taiwan University, Taipei, Taiwan; and the ³Department of Internal Medicine, National Taiwan University Hospital, Taipei, Taiwan.

Corresponding authors: Lee-Ming Chuang, leeming@ntu.edu.tw, and Juey-Jen Hwang, jueyhwang@ntu.edu.tw. Received 14 April 2009 and accepted 18 August 2009. Published ahead of print at http://care. diabetesjournals.org on 15 September 2009. DOI: 10.2337/dc09-0694.

J.-W.L. and Y.-C.C. contributed equally to this work.

© 2009 by the American Diabetes Association. Readers may use this article as long as the work is properly cited, the use is educational and not for profit, and the work is not altered. See http://creativecommons.org/licenses/by-nc-nd/3.0/ for details.

The costs of publication of this article were defrayed in part by the payment of page charges. This article must therefore be hereby marked "advertisement" in accordance with 18 U.S.C. Section 1734 solely to indicate this fact.

Table 1—The predictive performance of 10 diabetes risk scores for screening-detected diabetes, metabolic syndrome, and CKD

Model	ROC area (95% CI)	H-L <i>P</i> value	Sensitivity	Specificity	Correctly classified	LR ⁺	LR ⁻	Youden index	Number of predictors
Screening-detected									
diabetes									
ARIC (U.S.)	0.74 (0.70-0.77)	0.01	0.68	0.70	0.69	2.23	0.46	0.39	6
QDScore (U.K.)	0.74 (0.70-0.77)	< 0.0001	0.73	0.63	0.64	1.99	0.42	0.38	9
Cambridge (U.K.)	0.73 (0.69-0.76)	0.23	0.64	0.67	0.67	1.95	0.54	0.31	7
FINDRISC (Finnish)	0.73 (0.69-0.77)	0.78	0.67	0.67	0.67	2.01	0.49	0.34	8
Oman	0.72 (0.69-0.75)	0.08	0.65	0.67	0.67	1.99	0.52	0.32	5
Danish	0.72 (0.68-0.76)	0.07	0.63	0.70	0.69	2.08	0.58	0.33	6
Thai	0.71 (0.67-0.74)	0.04	0.71	0.62	0.62	1.87	0.47	0.33	6
Asian Indian	0.70 (0.66-0.74)	0.0005	0.63	0.69	0.68	2.01	0.54	0.32	5
Dutch	0.69 (0.64-0.73)	0.016	0.61	0.70	0.70	2.03	0.56	0.31	4
DESIR (French)	0.67 (0.64-0.71)	0.009	0.55	0.70	0.69	1.82	0.64	0.28	4
Metabolic syndrome									
DESIR (French)	0.82 (0.81-0.84)	0.002	0.70	0.83	0.79	4.06	0.37	0.53	4
Thai	0.82 (0.81-0.84)	< 0.0001	0.77	0.75	0.75	3.03	0.30	0.52	6
ARIC (U.S.)	0.81 (0.79-0.83)	0.62	0.74	0.73	0.73	2.72	0.35	0.47	6
Cambridge (U.K.)	0.80 (0.79-0.82)	< 0.0001	0.77	0.69	0.71	2.48	0.33	0.47	7
Asian Indian	0.78 (0.76-0.80)	< 0.0001	0.71	0.73	0.72	2.66	0.40	0.45	5
FINDRISC (Finnish)	0.77 (0.75-0.79)	< 0.0001	0.71	0.67	0.68	2.12	0.43	0.40	8
QDScore (U.K.)	0.77 (0.75-0.79)	< 0.0001	0.73	0.68	0.69	2.26	0.39	0.41	9
Danish	0.77 (0.76-0.79)	0.002	0.79	0.62	0.66	2.08	0.37	0.41	6
Dutch	0.73 (0.71-0.75)	< 0.0001	0.57	0.77	0.71	2.46	0.55	0.33	4
Oman	0.73 (0.71-0.74)	< 0.0001	0.76	0.62	0.66	2.02	0.39	0.41	5
CKD									
ARIC (U.S.)	0.71 (0.68-0.73)	0.46	0.64	0.68	0.67	1.97	0.53	0.31	6
Cambridge (U.K.)	0.68 (0.66-0.70)	0.002	0.69	0.55	0.58	1.54	0.56	0.25	7
QDScore (U.K.)	0.68 (0.65-0.70)	< 0.0001	0.64	0.61	0.61	1.64	0.59	0.28	9
Danish	0.67 (0.65-0.69)	< 0.0001	0.62	0.63	0.63	1.63	0.63	0.27	6
Dutch	0.66 (0.64-0.69)	< 0.0001	0.59	0.74	0.72	2.27	0.55	0.33	4
Oman	0.66 (0.64-0.69)	< 0.0001	0.58	0.65	0.63	1.64	0.65	0.22	5
Asian Indian	0.65 (0.62-0.67)	< 0.0001	0.63	0.59	0.60	1.53	0.63	0.24	5
FINDRISC (Finnish)	0.62 (0.59-0.64)	0.006	0.68	0.49	0.52	1.32	0.66	0.17	8
Thai	0.61 (0.58-0.63)	< 0.0001	0.56	0.59	0.59	1.38	0.74	0.16	6
DESIR (French)	0.60 (0.58-0.63)	0.01	0.77	0.40	0.47	1.28	0.58	0.17	4
T1 1	1.6. 1. 11.	1 1 . /1			7 1 1		1 1		C /

The optimal cutoff value was defined as the one with the least $(1 - \text{sensitivity})^2 + (1 - \text{specificity})^2$. Youden index was defined as the maximum of (sensitivity + specificity -1). H-L, Hosmer-Lemeshow goodness-of-fit test; LR⁺, positive likelihood ratio; LR⁻, negative likelihood ratio.

strongly correlate with DRSs (supplementary Table S1). All DRSs were associated with reduced MDRD GFR and with increased urine albumin excretion (supplementary Table S1). The prevalence of screening-detected diabetes (SDM), MetSyn, and CKD increased significantly with increasing DRSs (all *P* for trend <0.0001, supplementary Figs. S1–S3).

The predictive performance of 10 DRSs for SDM, MetSyn, and CKD are summarized in Table 1. The best area under the ROC for SDM was 0.74 (95% CI 0.70–0.77, Table 1), with 68% sensitivity and 70% specificity using optimal cutoff values. There were no statistical differences in the area under the ROC for SDM among ARIC, QDScore, Cambridge, FINDRISC, Oman, Danish, and Thai scores. How-

ever, the Cambridge risk score and the FINDRISC outperformed the other DRSs in model fit. Using stepwise logistic regression, we identified age, waist circumference, medication for hypertension, and family history of diabetes as independent predictors for SDM. The area under ROC of the logistic regression model based on these predictors was 0.75 (95% CI 0.71–0.78) for SDM, similar to those of DRSs.

The best area under the ROC for MetSyn was 0.82 (95% CI 0.81–0.84, Table 1) with 70% sensitivity and 83% specificity. There were no statistical differences in the areas under the ROC among the DESIR, Thai, and ARIC scores for MetSyn. However, the ARIC score outperformed other DRSs in model fit.

The best area under the ROC curves for CKD was 0.71 (95% CI 0.68–0.73) (supplementary Fig. S2). The sensitivity was 64% and specificity 68% to detect CKD. The ARIC score outperformed other DRSs in discriminative performance for CKD and model fit.

CONCLUSIONS

This study validated the predictive performance of currently available DRSs based on simple clinical information for type 2 diabetes in a cross-sectional screening program in Taiwan. The predictive performance of these DRSs among Taiwanese was comparable to those in other European populations (13). These data indicate that DRSs based on simple clinical information without laboratory test are

Diabetes risk scores in Taiwan

also applicable for cross-sectional screening across different ethnic populations. All DRSs strongly correlate with markers of insulin resistance and are strong predictors for MetSyn. Therefore, DRSs are actually predictors of insulin resistance, which in turn predicts type 2 diabetes.

Another key finding of this study was that all DRSs correlated with GFR and urine albumin excretion and are fair predictors of CKD. Although a great deal of effort has been exerted to develop a renal risk score to identify individuals at high risk of CKD, there is currently no simple and widely established renal risk score (14). Most proposed scoring models use serum creatinine and urine protein as the main predictors, which are often not available in large-scale population screening (14). Although the predictive performance of DRSs cannot surpass current renal risk models, the results here demonstrate the potential feasibility of developing a simple renal risk score to select high-risk individuals for further laboratory screening.

This study has some limitations. First, an oral glucose tolerance test was not performed and 2-h glucose concentration was not included in the definition of diabetes. Second, other DRSs including the American Diabetes Association risk tool, the Second National Health and Nutrition Survey questionnaire by Herman et al., the German risk score by Shulze et al., and the new ARIC risk scores by Kahn et al. were not included in the analyses because of the lack of delivery history, gestational diabetes history, detailed dietary information, and ethnic-specific cutoff values for weight and height (15-17).

Acknowledgments— This work was supported by grants from the National Taiwan

University Hospital Yun-Lin Branch, Taiwan (NTUHYL 98 X001 and NTUHYL 98 X008).

No potential conflicts of interest relevant to this article were reported.

We express our sincere gratitude to all participants.

References

- 1. Levey AS, Bosch JP, Lewis JB, Greene T, Rogers N, Roth D. A more accurate method to estimate glomerular filtration rate from serum creatinine: a new prediction equation: Modification of Diet in Renal Disease Study Group. Ann Intern Med 1999;130:461–470
- 2. Tan CE, Ma S, Wai D, Chew SK, Tai ES. Can we apply the national cholesterol education program adult treatment panel definition of the metabolic syndrome to Asians? Diabetes Care 2004;27:1182–1186
- Schmidt MI, Duncan BB, Bang H, Pankow JS, Ballantyne CM, Golden SH, Folsom AR, Chambless LE. The Atherosclerosis Risk in Communities Investigators: identifying individuals at high risk for diabetes: the Atherosclerosis Risk in Communities Study. Diabetes Care 2005;28:2013– 2018
- Ramachandran A, Snehalatha C, Vijay V, Wareham NJ, Colagiuri S. Derivation and validation of diabetes risk score for urban Asian Indians. Diabetes Res Clin Pract 2005;70:63–70
- Griffin SJ, Little PS, Hales CN, Kinmonth AL, Wareham NJ. Diabetes risk score: towards earlier detection of type 2 diabetes in general practice. Diabete Metab Res Rev 2000;16:164–171
- Glümer C, Carstensen B, Sandbaek A, Lauritzen T, Jørgensen T, Borch-Johnsen K. A Danish diabetes risk score for targeted screening: the Inter99 study. Diabetes Care 2004;27:727–733
- 7. Balkau B, Lange C, Fezeu L, Tichet J, de Lauzon-Guillain B, Czernichow S, Fumeron F, Froguel P, Vaxillaire M, Cauchi S, Ducimetière P, Eschwège E. Predicting diabetes: clinical, biological, and genetic approaches: data from the Epidemiological Study on the Insulin Resistance Syndrome

- (DESIR). Diabetes Care 2008;31:2056-2061
- 8. Baan CA, Ruige JB, Stolk RP, Witteman JC, Dekker JM, Heine RJ, Feskens EJ. Performance of a predictive model to identify undiagnosed diabetes in a health care setting. Diabetes Care 1999;22:213–219
- Lindström J, Tuomilehto J. The diabetes risk score: a practical tool to predict type 2 diabetes risk. Diabetes Care 2003;26:725– 731
- Al-Lawati JA, Tuomilehto J. Diabetes risk score in Oman: a tool to identify prevalent type 2 diabetes among Arabs of the Middle East. Diabetes Res Clin Pract 2007;77: 438–444
- Hippisley-Cox J, Coupland C, Robson J, Sheikh A, Brindle P. Predicting risk of type 2 diabetes in England and Wales: prospective derivation and validation of QDScore. BMJ 2009;338:b880
- Aekplakorn W, Bunnag P, Woodward M, Sritara P, Cheepudomwit S, Yamwong S, Yipintsoi T, Rajatanavin R. A risk score for predicting incident diabetes in the Thai population. Diabetes Care 2006;29:1872– 1877
- 13. Schwarz PEH, Li J, Lindstrom J, Tuimilehto J. Tools for predicting the risk of type 2 diabetes in daily practice. Horm Metab Res 2009;41:86–97
- Taal MW, Brenner BM. Renal risk scores: progress and prospects. Kidney Int 2008; 73:1216–1219
- 15. Herman WH, Smith PJ, Thompson TJ, Engelgau MM, Aubert RE. A new and simple questionnaire to identify people at increased risk for undiagnosed diabetes. Diabetes Care 1995;18:382–387
- 16. Schulze MB, Hoffmann K, Boeing H, Linseisen J, Rohrmann S, Möhlig M, Pfeiffer AF, Spranger J, Thamer C, Häring HU, Fritsche A, Joost HG. An accurate risk score based on anthropometric, dietary, and lifestyle factors to predict the development of type 2 diabetes. Diabetes Care 2007;30:510–515
- 17. Kahn HS, Cheng YJ, Thompson TJ, Imperatore G, Gregg EW. Two risk-scoring systems for predicting incident diabetes mellitus in U.S. adults age 45 to 64 years. Ann Intern Med 2009;150:741–751